Pharmacy Policy Bulletin

Title: Heart Failure Agents
Policy #: Rx.01.174

Application of pharmacy policy is determined by benefits and contracts. Benefits may vary based on product line, group, or contract. Some medications may be subject to precertification, age, gender or quantity restrictions. Individual member benefits must be verified.

This pharmacy policy document describes the status of pharmaceutical information and/or technology at the time the document was developed. Since that time, new information relating to drug efficacy, interactions, contraindications, dosage, administration routes, safety, or FDA approval may have changed. This Pharmacy Policy will be regularly updated as scientific and medical literature becomes available. This information may include new FDA-approved indications, withdrawals, or other FDA alerts. This type of information is relevant not only when considering whether this policy should be updated, but also when applying it to current requests for coverage.

Members are advised to use participating pharmacies in order to receive the highest level of benefits.

Intent:
The intent of this policy is to communicate the medical necessity criteria for Corlanor® (ivabradine) and Entresto™ (sacubitril/valsartan) as provided under the member's pharmacy benefit.

Description:
Heart failure affects approximately 5.1 million Americans. While survival after a diagnosis of heart failure has improved, it is still about 50% at 5 years after diagnosis. In patients with symptomatic heart failure and left ventricular ejection fraction (LVEF) < 40%, current guidelines recommend angiotensin converting enzyme (ACE) inhibitors or angiotensin receptor blockers (ARB), beta blockers, diuretics and aldosterone antagonists as standard of care. With the exception of diuretics, these classes of medications decrease mortality from heart failure with reduced EF. Diuretics provide symptomatic relief, however the effects of diuretics on morbidity and mortality are not known.

Ivabradine reduces spontaneous pacemaker activity at the cardiac sinus node by blocking the hyperpolarization-activated cyclic nucleotide-gated (HCN) channel to selectively inhibit I$_f$-current, thus reducing the heart rate. Ventricular repolarization and myocardial contractility are not affected. Increased heart rate is an ineffective compensatory mechanism and is recognized as an independent risk factor in heart failure.

Corlanor (ivabradine) is indicated to reduce the risk of hospitalization for worsening heart failure in patients with stable, symptomatic chronic heart failure with left ventricular ejection fraction ≤ 35%, who are in sinus rhythm with resting heart rate ≥ 70 beats per minute and either are on maximally tolerated doses of beta blockers or have a contraindication to beta-blocker use.

Sacubitril inhibits nephrilysin via its active metabolite, LBQ657. Valsartan inhibits the effects angiotensin
(AT) II by selectively blocking the AT1 receptor.

Entresto (sacubitril/valsartan) is indicated to reduce the risk of cardiovascular death and hospitalization for heart failure in patients with chronic heart failure (NYHA Class II-IV) and reduced ejection fraction. Entresto is usually administered in conjunction with other heart failure therapies, in place of an ACE inhibitor or other ARB.

Policy:

Corlanor (ivabradine) is approved when ALL of the following are met:

1. Stable, symptomatic chronic heart failure; and
2. LVEF ≤ 35%; and
3. Sinus rhythm with resting heart rate ≥ 70 beats per minute; and
4. Patient is clinically stable for at least 4 weeks on an optimized and stable clinical regimen which includes:
   a. Maximally tolerated doses of beta blockers or inability to tolerate beta blockers; and
   b. ACE inhibitors or ARBs or inability to tolerate ACE inhibitor or ARB

Entresto (sacubitril/valsartan) is approved when ALL of the following are met:

1. Member is 18 years of age or older; and
2. Diagnosis of chronic heart failure with NYHA functional class II-IV; and
3. Ejection fraction less than 35%; and
4. Concurrent administration of or inability to tolerate beta blockers

Black Box Warning:

Entresto (sacubitril/valsartan): When pregnancy is detected, discontinue ENTRESTO as soon as possible. Drugs that act directly on the renin-angiotensin system can cause injury and death to the developing fetus.

Guidelines:

Refer to the specific manufacturer's prescribing information for administration and dosage details and any applicable Black Box warnings.

BENEFIT APPLICATION

Subject to the terms and conditions of the applicable benefit contract, the applicable drug(s) identified in this policy is (are) covered under the pharmacy benefits of the Company’s products when the medical necessity criteria listed in this pharmacy policy are met. Any services that are experimental/investigational or cosmetic are benefit contract exclusions for all products of the Company.

References:


Applicable Drugs:

<table>
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<th>Brand Name</th>
<th>Generic Name</th>
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<tr>
<td>Corlanor</td>
<td>Ivabradine</td>
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<td>Entresto</td>
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Cross References:

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